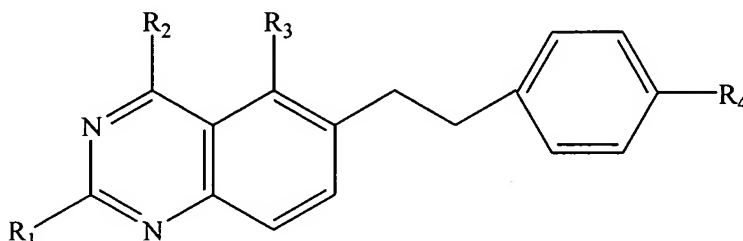


Listing of the Claims:

1. and 2. (Canceled)

3. (Previously Presented) In a process for synthesizing a compound of formula Ib:

(Ib)



wherein R₁ and R₂ are each individually amino or N-alkyl substituted amino; hydroxy; alkoxy; keto; lower alkyl;

R₃ is hydrogen; hydroxy; alkoxy; trifluoromethyl alkoxy; halo; sulfhydryl or alkylthio;

R₄ is -C(O)-X;

X is alkoxy; or γ -methylene glutamic acid, glutamic acid, aspartic acid or γ -methylene glutamate residue;

in which process a 2-amino-5-nitro-benzonitrile starting reagent is cyclized to form 2,4-diamino-6-nitro-quinazoline, which is converted to 2,4,6-triamino-quinazoline, which is converted to 2,4-diamino-6-cyano-quinazoline, which is converted to 2,4-diamino-6-formyl-quinazoline;

the improvement comprising:

reacting a bromo-alkyl-*p*-benzoic acid moiety with triethyl phosphite to form a 4-R₄-carbonyloxyalkyl-phenyl-alkyldiethylphosphonate; and

reacting the 2,4-diamino-6-formyl-quinazoline with the 4-R₄-carbonyloxyalkyl-phenyl-alkyldiethylphosphonate to form a product with an unsaturated bond; and

reducing said product to form the compound of formula Ib.

4. (Previously Presented) The process of claim 3, wherein the product with an unsaturated bond moiety is a methylene moiety.

5. (Previously Presented) The process of claim 4, wherein methylene moiety is 4-(1,2-methylene)benzoate.

6. (Canceled)

7. (Canceled)

8. (Previously Presented) The process of claim 3, wherein the 4-R₄-carbonyloxyalkyl-phenyl-alkyldiethylphosphonate is 4-carboxymethyl-phenyl-methyldiethylphosphonate.

9. (Previously Presented) The process of claim 3, further comprising hydrogenating the compound of formula Ib to form 6-(4-R₄-carbonyloxyalkylphenyl)ethanyl-2,5-diamino quinazoline, which is hydrolyzed to form 6-(4-R₄-carbonyloxyphenyl)ethanyl-2,5-diamino quinazoline,

the improvement further comprising:

reacting 6-(4-R₄-carbonyloxyphenyl)ethanyl-2,5-diamino quinazoline with diethyl γ -methylene-L-glutamate to form γ -methylene glutamate 5,8,10-trideaza aminopterin diethyl ester.

10. (Previously Presented) The process of claim 9, wherein the improvement further comprises reacting the 6-(4-R₄-carbonyloxyphenyl)ethanyl-2,5-diamino quinazoline with the diethyl γ -methylene-L-glutamate in the presence of 1-hydroxy benzotriazole and 1-[3-dimethylamino)propyl]-3-ethyl carbodiimide hydrochloride.

11. (Previously Presented) The process of claim 9, wherein the improvement further comprises reacting the 6-(4-R₄-carbonyloxyphenyl)ethanyl-2,5-diamino quinazoline with the diethyl γ -methylene-L-glutamate in the presence of 1-hydroxy benzotriazole, 1-[3-dimethylamino)propyl]-3-ethyl carbodiimide hydrochloride and triethylamine.